

Form PTO-1449 (modified)

List of Patents and Publications for Applicant

INFORMATION DISCLOSURE STATEMENT

(Use several sheets if necessary)

Atty. Docket No.

ILEX:040US/SLH

Serial No.

09/529,319

Applicant

Richard Poulin *et al.*

Filing Date:

April 11, 2000

Group:

Unknown 7625

U.S. Patent Documents

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Foreign Patent Documents

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Other Art

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U.S. Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Name	Class	Sub Class	Filing Date of App.
8e	A1	5,456,908	10/10/95	Aziz <i>et al.</i>	424	78.08	3/1/94

Foreign Patent Documents

Exam. Init.	Ref. Des.	Document Number	Date	Country	Class	Sub Class	Translation Yes/No
8e	B1	WO 93/12777	7/8/93	PCT			
8e	B2	WO 98/17623	4/30/98	PCT			

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Exam. Init.	Ref. Des.	Citation
8e	C1	Ask <i>et al.</i> , "Antileukemic effects of non-metabolizable derivatives of spermidine and spermine," <i>Cancer Lett.</i> , 69:33-38, 1993.
	C2	Ask <i>et al.</i> , "Increased survival of L1210 leukemic mice by prevention of the utilization of extracellular polyamines. Studies using a polyamine-uptake mutant, antibiotics and a polyamine-deficient diet," <i>Cancer Lett.</i> , 66:29-34, 1992.
	C3	Aziz <i>et al.</i> , "A novel polymeric spermine conjugate inhibits polyamine transport in pulmonary artery smooth muscle cells," <i>J. Pharmacol. Exper. Ther.</i> , 274:181-186, 1992.
	C4	Aziz <i>et al.</i> , "The potential of a novel polyamine transport inhibitor in cancer chemotherapy," <i>Pharmacol. Exper. Ther.</i> , 278:185-192, 1996.
	C5	Bergeron <i>et al.</i> , "Development of a hypusine reagent for peptide synthesis," <i>Org. Chem.</i> , 62:3285-3290, 1997.
	C6	Chaney <i>et al.</i> , "Tumor selective enhancement of radioactivity uptake in mice treated with α -difluoromethylornithine prior to administration of ^{14}C -putrescine," <i>Life Sci.</i> , 32:1237-1241, 1983.
	C7	Chang <i>et al.</i> , "Modulation of polyamine biosynthesis and transport by oncogene transfection," <i>Biochem. Biophys. Res. Comm.</i> , 157:264-270, 1988.

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EXAMINER:

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List of Patents and Publications for Applicant's INFORMATION DISCLOSURE STATEMENT (Use several sheets if necessary)		Applicant Richard Poulin <i>et al.</i>	
		Filing Date: April 11, 2000	Group: Unknown 162)
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82	C8	Cohen <i>et al.</i> , "Targeting of cytotoxic agents by polyamines: synthesis of a chlorambucil-spermidine conjugate," <i>J. Chem. Soc. Chem. Commun.</i> , pp. 298-300, 1992.
	C9	Duranton <i>et al.</i> , "Suppression of preneoplastic changes in the intestine of rats fed low levels of polyamines," <i>Cancer Res.</i> , 57:573-575, 1997.
	C10	Felschow <i>et al.</i> , "Photoaffinity labeling of a cell surface polyamine binding protein," 270:28705-28711, 1995.
	C11	Frebort and Adachi, "Copper/quinone-containing amine oxidases, an exciting class of ubiquitous enzymes," <i>J. Ferment. Bioeng.</i> , 80:625-632, 1995.
	C12	Hayashi <i>et al.</i> , Ornithine decarboxylase antizyme - A novel type of regulatory protein," <i>Trends Biochem. Sci.</i> , 21:27-30, 1996.
	C13	He <i>et al.</i> , "Antizyme delays the restoration by spermine of growth of polyamine-deficient cells through its negative regulation of polyamine transport," <i>Biochem. Biophys. Res. Commun.</i> , 203:608-614, 1994.
	C14	Hessels <i>et al.</i> , "Microbial flora in the gastrointestinal tract abolishes cytostatic effects of α -difluoromethylornithine in vivo," <i>Int. J. Cancer</i> , 43:1155-1164, 1989.
	C15	Holley <i>et al.</i> , "Targeting of tumor cells and DNA by a chlorambucil-spermidine conjugate," <i>Cancer Res.</i> , 52:4190-4195, 1992.
	C16	Horn, <i>et al.</i> , "Phase I-II clinical trial with α -difluoromethylornithine- an inhibitor of polyamine biosyntheses," <i>Eur. J. Cancer Clin. Oncol.</i> , 23:1103-1107, 1987.
	C17	Huber and Poulin, "Antiproliferative effect of spermine depletion by -cyclohexyl-1,3-diaminopropane in human breast cancer cell growth," <i>Cancer Res.</i> , 55:934-943, 1995.
	C18	Huber and Poulin, "Permissive role of polyamines in the cooperative action of estrogens and insulin or insulin-like growth factor I on human breast cancer cell growth," <i>J. Clin. Endocrinol. Metab.</i> , 81:113-123, 1996.
	C19	Huber <i>et al.</i> , "2,2'-Dithiobis(N-ethyl-spermine-5-carboximide) is a high affinity, membrane-impermeant antagonist of the mammalian polyamine transport system," <i>J. Biol Chem.</i> , 271:27556-27563, 1996.
	C20	Janne <i>et al.</i> , "Polyamines from molecular biology to clinical applications," <i>Ann. Med.</i> , 23:241-259, 1991.

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82	C21	Kanter <i>et al.</i> , "Preclinical toxicologic evaluation of DENSPM (N ¹ , N ¹¹ -diethylnorspermine) in rats and dogs," <i>Anti-Cancer Drugs</i> , 5:448-456, 1994.
	C22	Lakanen <i>et al.</i> , "α-Methyl polyamines: metabolically stable spermidine and spermine mimics capable of supporting growth in cells depleted of polyamines," <i>J. Med. Chem.</i> , 35:724-734, 1992.
	C23	Lessard <i>et al.</i> , "Hormonal and feedback regulation of putrescine and spermidine transport in human breast cancer cells," <i>J. Biol. Chem.</i> 270:1685-1694, 1995.
	C24	Li <i>et al.</i> , "Comparative molecular field analysis-based predictive model of structure-function relationships of polyamine transport inhibitors in L1210 cells," <i>Cancer Res.</i> , 57:234-239, 1997.
	C25	Love <i>et al.</i> , "Randomized phase I chemoprevention dose-seeking study of α-difluoromethylornithine," <i>J. Natl. Cancer Inst.</i> , 85:732-737, 1993.
	C26	Marton and Pegg, "Polyamines as targets for therapeutic intervention," <i>Ann. Rev. Pharmacol. Toxicol.</i> , 35:55-91, 1995.
	C27	Mausumoto and Suzuki, "Polyamines as markers of malignancy," In: <i>The Physiology of Polyamines</i> , edited by U. Bachrach and Y.M. Heimer. Boca Raton, FL: CRC Press, 219-234, 1989.
	C28	McCann and Bitonti, "An overview of inhibition of polyamine metabolism and the consequent effects on cell proliferation in mammalian cells and parasitic protozoa," <i>Polyamines in the Gastrointestinal Tract</i> , edited by R.H. Dowling, U.R. Folsch and C. Loser, Dordrecht: Kluwer Academic Publ., 143-153, 1992.
	C29	McCann and Pegg, "Ornithine decarboxylase as an enzyme target for therapy," <i>Pharmac. Ther.</i> , 54:195-215, 1992.
	C30	McCormack and Johnson, "Putrescine uptake and release by colon cancer cells," <i>Am. J. Physiol.</i> , 256:G868-G877, 1989.
	C31	Meyskens and Gerner, "Development of difluoromethyl-ornithine as chemoprevention agent for the management of colon cancer," <i>J. Cell. Biochem.</i> , 22:126-131, 1995.
2	C32	Minchin <i>et al.</i> , "Inhibition of putrescine uptake by polypyridinium quaternary salts in B16 melanoma cells treated with difluoromethylornithine," <i>Biochem. J.</i> , 262:391-395, 1989.

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gle	C33	Mitchell <i>et al.</i> , "Feedback repression of polyamine transport is mediated by antizyme in mammalian tissue culture cells," <i>Biochem. J.</i> , 299:19-22, 1994.
	C34	Mitchell <i>et al.</i> , "Feedback repression of polyamine uptake into mammalian cells require active protein synthesis," <i>Biochem Biophys. Res. Commun.</i> , 186:81-88, 1992.
	C35	Morgan, "Polyamine oxidases and oxidized polyamines," In: <i>The Physiology of Polyamines</i> , edited by U. Bachrach and U.M. Heimer. Boca Raton, FL: CRC Press, p. 203-229, 1989.
	C36	Moulinoux <i>et al.</i> , "Biological significance of circulating polyamines in oncology," <i>Cell. Mol. Biol.</i> , 37:773-783, 1991.
	C37	Moulinoux <i>et al.</i> , "The growth of MAT-LyLu rat prostatic adeno-carcinoma can be prevented in vivo by polyamine deprivation," <i>J. Urol.</i> , 146:1408-1412, 1991.
	C38	Nicolet <i>et al.</i> , "Putrescine and spermidine uptake is regulated by proliferation and dexamethasone treatment in AR4-2J cells," <i>Int. J. Cancer</i> , 49:577-581, 1991.
	C39	O'Sullivan <i>et al.</i> , "Inhibiting effects of spermidine derivatives on <i>Trypanosoma cruzi</i> trypanothione reductase," <i>J. Enzym Inhib.</i> , 11:97-114, 1996.
	C40	Osborne and Seidel, "Gastrointestinal luminal polyamines: cellular accumulation and enterohepatic circulation," <i>Am. J. Physiol.</i> , 258:G576-G584, 1990.
	C41	Parchment <i>et al.</i> , "Serum amine oxidase activity contributes to crisis in mouse embryo cell lines," <i>Proc. Natl. Acad. Sci., USA</i> 87:4340-4344, 1990.
	C42	Pegg <i>et al.</i> , "Effect of S-adenosyl-1, 12-diamino-3-thio-9-azadodecane, a multisubstrate adduct inhibitor of spermine synthase, on polyamine metabolism in mammalian cells," <i>Biochemistry</i> , 28:8446-8453, 1989.
	C43	Pegg <i>et al.</i> , "Inhibition of polyamine biosynthesis and function as an approach to drug design," <i>Enzymes as Targets for Drug Design</i> , edited by M.G. Palfreyman, P.P. McCann, P.P. Lovenberg, W. Temple, J.G. Temple and A. Sjoerdsma. Orlando: Academic Press, 157-183, 1989.
	C44	Pegg <i>et al.</i> , "Use of aminopropyltransferase inhibitors and of non-metabolizable analogs to study polyamine regulation and function," <i>Int. J. Biochem. Cell. Biol.</i> , 27:425-442, 1995.
u	C45	Persson <i>et al.</i> , "Curative effect of DL-2 difluoromethylornithine on mice bearing mutant L1210 leukemia cells deficient in polyamine uptake," <i>Cancer Res.</i> , 48:4807-4811, 1988.

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812	C46	Porter and Bergeron, "Spermidine requirement for cell proliferation in eukaryotic cells: structural specificity and quantitation," <i>Science</i> , 219:1083-1085, 1983.
	C47	Porter <i>et al.</i> , "Antitumor activity of N ¹ , N ¹¹ -bis(ethyl)norspermine against human melanoma xenografts and possible biochemical correlates of drug action," <i>Cancer Res.</i> , 53:581-586, 1993.
	C48	Porter <i>et al.</i> , "Biological properties of N ⁴ spermidine derivatives and their potential in anticancer chemotherapy," <i>Cancer Res.</i> , 42:4072-4078, 1982.
	C49	Porter <i>et al.</i> , "Biological properties of N ⁴ - and N ¹ , N ⁸ -spermidine derivatives in cultured L1210 leukemia cells," <i>Cancer Res.</i> , 45:2050-2057, 1985.
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	C53	Pusztai <i>et al.</i> , "Stimulation of growth and polyamine accretion in the small intestine and pancreas by lectins and trypsin inhibitors," <i>Polyamines in the Gastrointestinal Tract</i> , edited by R.H. Dowling, U.R. Folsch and C. Loser. Dordrecht:Kluwer Academic Publ., 473-483, 1992.
	C54	Quemener <i>et al.</i> , "Polyamine deprivation: a new tool in cancer treatment," <i>Anticancer. Res.</i> , 14:443-448, 1994.
	C55	Quemener <i>et al.</i> , "Tumour growth inhibition by polyamine deprivation," <i>Polyamines in the Gastrointestinal Tract</i> , edited by R.H. Dowling, U.R. Folsch and C. Loser. Dordrecht:Kluwer Academic Publ., 375-385, 1992.
	C56	Rinehart and Chen, "Characterization of the polyamine transport system in mouse neuroblastoma cells," <i>J. Biol. Chem.</i> , 259:4750-4756, 1984.
↓	C57	Sarhan <i>et al.</i> , "The gastrointestinal tract as polyamine source for tumor growth," <i>Anticancer Res.</i> , 9:215-224, 1989.

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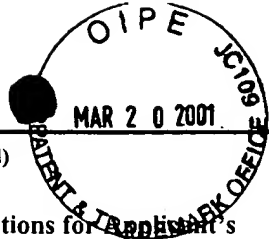
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SL	C58	Schechter <i>et al.</i> , "Clinical aspects of inhibition of ornithine decarboxylase with emphasis on therapeutic trials of eflornithine (DFMIO) in cancer and protozoan diseases," <i>Inhibition of Polyamine Metabolism. Biological Significance and Basis for New Therapies</i> , edited by P.P. McCann, A.E. Pegg and A. Sjoerdsma. Orlando, FL:Academic Press, 345-364, 1987.
	C59	Seiler and Dezeure, "Polyamine transport in mammalian cells," <i>Int J. Biochem.</i> , 22:211-218, 1990.
	C60	Seiler <i>et al.</i> , "Endogenous and exogenous polyamines in support of tumor growth," <i>Cancer Res.</i> , 50:5077-5083, 1990.
	C61	Seiler <i>et al.</i> , "Polyamine transport in mammalian cells. An update," <i>Int. J. Biochem. Cell Biol.</i> , 28:843-861, 1996.
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	C63	Sjoerdsma and Schechter, "Chemotherapeutic implication so polyamine biosynthesis inhibition," <i>Clin. Pharm. Thera.</i> , 35:287-300, 1984.
	C64	Sjoerdsma and Schechter, "Successful treatment of lethal protozoal infections with the ornithine decarboxylase inhibitor, α -difluoromethylornithine," <i>Trans. Ass. Am. Physic.</i> , 97:70-79, 1984.
	C65	Stark <i>et al.</i> , "Synthesis and evaluation of novel spermidine derivatives as targeted cancer chemotherapeutic agents," <i>J. Med. Chem.</i> , 35, 4264-4269, 1992.
	C66	Steele <i>et al.</i> , "Preclinical efficacy evaluation of potential chemo-preventive agents in animal carcinogenesis: methods and results from the NCI Chemoprevention Drug Development Program," <i>J. Cell. Biochem. (suppl.)</i> 20:32-54, 1994.
	C67	Sunkara, P. S. <i>et al.</i> , "Inhibitors of polyamine biosynthesis: cellular and <i>in vivo</i> effects on tumor proliferation," <i>Inhibition of Polyamine Metabolism Biological Significance and Basis for New Therapies.</i> , edited by P.P. McCann, A.E. Pegg and A. Sjoerdsma. Orlando:Academic Press, 121-140, 1987.
2	C68	Talpaz <i>et al.</i> , "Clinical studies of α -difluoromethylornithine and α -interferon combination in cancer patients," <i>The Physiology of Polyamines</i> , edited by U. Bachrach and Y.M. Heimer, Boca Raton, FL:CRC Press, 287-292, 1989.

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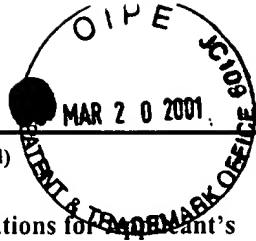
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80	C69	Tanaka <i>et al.</i> , "Chemoprevention of oral carcinogenesis by DL- α -difluoromethylornithine, an ornithine decarboxylase inhibitor: dose-dependent reduction in 4-nitroquinoline 1-oxide-induced tongue neoplasms in rats," <i>Cancer Res.</i> , 53:772-776, 1993.
	C70	Tempero <i>et al.</i> , "Chemoprevention of mouse colon tumors with difluoromethylornithine during and after carcinogen treatment," <i>Cancer Res.</i> , 49:5793-5797, 1989.
	C71	Tjandrawinata and Byus, "Regulation of the efflux of putrescine and cadaverine from rapidly growing cultured RAW 264 cells by extracellular putrescine," <i>Biochem. J.</i> , 305:291-299, 1995.
	C72	Tjandrawinata <i>et al.</i> , "Regulation of putrescine export in lipopolysaccharide or IFN- γ -activated murine monocytic-leukemic RAW 264 Cells," <i>J. Immunol.</i> , 152:3039-3052, 1994.
2	C73	Zang and Sadler, "Synthesis of hexamine ligands by using trityl as an N-blocking agent," <i>Synthetic Communications</i> , 27:3145-3150, 1997.

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